

## RAW SEQUENCE LISTING ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number

Date Processed by STIC:

10 658,355pt T+WD; 3/16/54

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.
PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR CRF SUBMISSION AND PATENTIN SOFTWARE QUESTIONS, PLEASE CONTACT MARK SPENCER, TELEPHONE: 703-308-4212; FAX: 703-308-4221 <u>Effective 12/13/03</u>: TELEPHONE: 571-272-2510; FAX: 571-273-0221

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE <u>CHECKER VERSION 4.1 PROGRAM</u>, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS.

http://www.uspto.gov/web/offices/pac/checker/chkr41note.htm

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail. Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom. Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses.

- 1. EFS-Bio (<a href="http://www.uspto.gov/ebc/efs/downloads/documents.htm">http://www.uspto.gov/ebc/efs/downloads/documents.htm</a>, EFS Submission User Manual ePAVE)
- 2. U.S. Postal Service: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450
- 3. Hand Carry directly to (EFFECTIVE 12/01/03):
  U.S. Patent and Trademark Office, Box Sequence, Customer Window, Lobby, Room 1B03, Crystal Plaza Two.
  2011 South Clark Place, Arlington, VA 22202
- 4. Federal Express, United Parcel Service, or other delivery service to: U.S. Patent and Trademark Office. Box Sequence, Room 4D03-Mailroom, Crystal Plaza Two, 2013 South Clark Place, Arlington, VA 22202

Revised 10/08/03

## Raw Sequence Listing Error Summary

-	RROR DETECTED	SUGGESTED CORRECTION SERIAL NUMBER: 101658, 355
A	TTN: NEW RULES CASE	S: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE
	I V Wrapped Nucleics Wrapped Aminos	The number/text at the end of each line "wrapped" down to the pout line. The
2	2Invalid Line Lengt	h The rules require that a line not exceed 72 characters in length. This includes white spaces.
3	Misaligned Amino Numbering	The numbering under each 5 <sup>th</sup> amino acid is misaligned. Do not use tab codes between numbers; use space characters, instead.
4	Non-ASCII	The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.
5	Variable Length	Sequence(s)contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.
6	PatentIn 2.0 "bug"	A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s) Normally, PatentIn would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.
7_	Skipped Sequences (OLD RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence:  (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown)  (i) SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading)  (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown)  This sequence is intentionally skipped
		Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to include the skipped sequences.
8_	Skipped Sequences (NEW RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence. <210> sequence id number <400> sequence id number 000
9	(NEW RULES)	Use of n's and/or Xaa's have been detected in the Sequence Listing.  Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present.  In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.
10_	Invalid <213> Response	Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence
11_		Sequence(s) missing the <220> "Feature" and associated numeric identifiers and responses.  Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section.  (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32) (Sec. 1.823 of Sequence Rules)
12_	PatentIn 2.0 "bug"	Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.
13		'n" can only represent a single nuclcotide; "Xaa" can only represent a single amino acid

AMC – Biotechnology Systems Branch – 09/09/2003



**IFWO** 

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/658,355A

DATE: 03/16/2004 TIME: 07:58:57

Input Set : D:\923seq.002

Output Set: N:\CRF4\03162004\J658355A.raw

```
3 <110> APPLICANT: Gantier, Rene
                     Guyon, Thierry
                      Hugo, Cruz Ramos
                      Vega, Manuel
                     Drittanti, Lila
         9 <120> TITLE OF INVENTION: Rational Directed Protein Evolution Using Two Dimensional
        10
                     Rational
                     Mutagenesis Scanning
        11
        13 <130> FILE REFERENCE: 38751-923
        15 <140> CURRENT APPLICATION NUMBER: US/10/658,355A
        16 <141> CURRENT FILING DATE: 2003-09-08
                                                                                                  Does Not Comply
        18 <150> PRIOR APPLICATION NUMBER: 60/457,063
                                                                                                  Correspon Tookatta Medale.
        19 <151> PRIOR FILING DATE: 2003-03-21
        21 <150> PRIOR APPLICATION NUMBER: 60/410,258
DNA

ORGANISM: Artificial Sequence

A 50 (220) FEATURE:

7451 (223) OTHER INFORMATION: EcoRI Forward Primer

7453 (400) SEQUENCE: 182

E--> 7454 geotgtatga tttattggat gttggaattc cctgatgegg tattttctcc ttacg

7455 (55)

7457 (210) SEQ ID NO: 183

T458 (211) LENGTH: 55

7459 (212) TYPE: DNA

7460 (213) ORGANISM: Artificial Sequence

7462 (220) FEATURE:

7463 (223) OTHER INFORMATION: EcoRI Reverse Primer

7465 (400) SEQUENCE: 183

-> 7466 cctaaggaga aaataccgca tcagggaatt

7467 (55)

7469 (210) SEQ ID NO: 104

7470 (211) LENGTH:
        22 <151> PRIOR FILING DATE: 2002-09-09
          7471 <212> TYPE: DNA
          7472 <213> ORGANISM: Artificial Sequence
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RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/658,355A

DATE: 03/16/2004 TIME: 07:58:58

Input Set : D:\923seq.002

Output Set: N:\CRF4\03162004\J658355A.raw

SAME Errors

```
7474 <220> FEATURE:
    7475 <223> OTHER INFORMATION: Seq ClaI Forward Primer
     7477 <400> SEQUENCE: 184
E--> 7478 ctgattatca accgcggtac atatgattga catgc
     7479 (35)
     7481 <210> SEQ ID NO: 185
     7482 <211> LENGTH: 31
     7483 <212> TYPE: DNA
     7484 <213> ORGANISM: Artificial Sequence
     7486 <220> FEATURE:
     7487 <223> OTHER INFORMATION: Seq ClaI Reverse Primer
     7489 <400> SEQUENCE: 185
E--> 7490 taogggataa taccgcgcca catagcagaa c
     7491 (31)
     7493 <210> SEQ ID NO: 186
     7494 <211> LENGTH: 18
     7495 <212> TYPE: DNA
     7496 <213> ORGANISM: Artificial Sequence
     7498 <220> FEATURE:
     7499 <223> OTHER INFORMATION: Seq Forward Primer
     7501 <400> SEQUENCE: 186
E--> 7502 cctgatgaag gaggactc
     7503 (18)
7505 <210> SEQ ID NO: 187
     7506 <211> LENGTH: 18
     7507 <212> TYPE: DNA
     7508 <213> ORGANISM: Artificial Sequence
      7510 <220> FEATURE:
     7511 <223> OTHER INFORMATION: Seq Reverse Primer
      7513 <400> SEQUENCE: 187
E--> 7514 ccaagcagca gatgagte
      7515 (18)--
     7517 <2TO> SEQ ID NO: 188
      7518 <211> LENGTH: 31
      7519 <212> TYPE: DNA
      7520 <213> ORGANISM: Artificial Sequence
      7522 <220> FEATURE:
      7523 <223> OTHER INFORMATION: IFN alpha-2b 5' Primer
      7525 <400> SEQUENCE: 188
 E--> 7526 tcagctgcaa gtcaagctgc tctgtgggct g
      7527 (31)
      7529 <2TO> SEQ ID NO: 189
      7530 <211> LENGTH: 48
      7531 <212> TYPE: DNA
      7532 <213> ORGANISM: Artificial Sequence
      7534 <220> FEATURE:
      7535 <223> OTHER INFORMATION: IFN alpha-2b 3' Primer
      7537 <400> SEQUENCE: 189
 E--> 7538 gctctagatc attecttact tettaaactt tettgeaagt ttgttgac
```

DATE: 03/16/2004

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TIME: 07:58:58
                    PATENT APPLICATION: US/10/658,355A
                                                                 same errors
                    Input Set : D:\923seq.002
                    Output Set: N:\CRF4\03162004\J658355A.raw
    7539 48
    7541 <210> SEQ ID NO: 190
    7542 <211> LENGTH: 36
    7543 <212> TYPE: DNA
    7544 <213> ORGANISM: Artificial Sequence
    7546 <220> FEATURE:
    7547 <223> OTHER INFORMATION: IFN alpha-2b HindIII Primer
                                                                            36
    7549 <400> SEQUENCE: 190
E--> 7550 ccgaagetta tggcettgae etttgettta etggtg
    7551 (36)---
    7553 <210> SEQ ID NO: 191
     7554 <211> LENGTH: 48
     7555 <212> TYPE: DNA
     7556 <213> ORGANISM: Artificial Sequence
     7558 <220> FEATURE:
     7559 <223> OTHER INFORMATION: IFN alpha-2b XbaI Primer
     7561 <400> SEQUENCE: 191
E--> 7562 getetagate attecttact tettaaactt tettgeaagt ttgttgac
                                                                         48
     7563 (48)
     7565 <210> SEQ ID NO: 192
     7566 <211> LENGTH: 80
     7567 <212> TYPE: DNA
     7568 <213> ORGANISM: Artificial Sequence
     7570 <220> FEATURE:
     7571 <223> OTHER INFORMATION: IFN alpha-2b 80 bp 5' Primer
     7573 <400> SEQUENCE: 192
E--> 7574 cccaagetta tggcettgae etttgettta etggtggece teetggtget eagetgeaag > 60
E--> 7575(80)
E--> 7576 tcaagctgct ctgtgggctg
     7578 <210> SEQ ID NO: 193
     7579 <211> LENGTH: 20
     7580 <212> TYPE: DNA
     7581 <213> ORGANISM: Artificial Sequence
     7583 <220> FEATURE:
     7584 <223> OTHER INFORMATION: EMCV Forward Primer
     7586 <400> SEQUENCE: 193
E--> 7587 cccctacatt gaggcatcca
     7588 (20)
     7590 <210> SEQ ID NO: 194
     7591 <211> LENGTH: 21
     7592 <212> TYPE: DNA
     7593 <213> ORGANISM: Artificial Sequence
     7595 <220> FEATURE:
     7596 <223> OTHER INFORMATION: EMCV Reverse Primer
     7598 <400> SEQUENCE: 194
E--> 7599 caggagcagg acaaggtcac t
     7600 (21)
     7602 <210> SEQ ID NO: 195
     7603 <211> LENGTH: 22
```

RAW SEQUENCE LISTING

The state of the s

DATE: 03/16/2004

TIME: 07:58:58

```
SAME
                     Input Set : D:\923seq.002
                     Output Set: N:\CRF4\03162004\J658355A.raw
     7604 <212> TYPE: DNA
     7605 <213> ORGANISM: Artificial Sequence
     7607 <220> FEATURE:
     7608 <221> NAME/KEY: misc feature
     7609 <222> LOCATION: 1; 22
     7610 <223> OTHER INFORMATION: EMCV Probe: n at position 1 is a 5-carboxyfluorescein
               (FAM) -modified deoxycytidylate; n at position 22 is an N, N, N', N'-
                tetramethyl-6-carboxyrhodamine (TAMRA)-modified deoxythymidylate.
     7612
     7614 <400> SEQUENCE: 195
E--> 7615 nagecgtca agacccaacc gen
     7616 (22)
     8074 <210> SEQ ID NO: 208
     8075 <211> LENGTH: 41
     8076 <212> TYPE: DNA
     8077 <213> ORGANISM: Artificial Sequence
     8079 <220> FEATURE:
     8080 <223> OTHER INFORMATION: Oligonucleotide
     8082 <400> SEQUENCE: 208
E--> 8083 aacatatgtg tgatctgcct caaacccaca gcctgggtag c
     8084 (41)
8086 <210> SEQ ID NO: 209
     8087 <211> LENGTH: 46
     8088 <212> TYPE: DNA
     8089 <213> ORGANISM: Artificial Sequence
     8091 <220> FEATURE:
     8092 <223> OTHER INFORMATION: Oligonucleotide
     8094 <400> SEQUENCE: 209
E--> 8095 aggatecte attecttact tettaaaett tettgeaagt ttgttg
     8096 (46)
     8098 <210> SEQ ID NO: 210
     8099 <211> LENGTH: 41
     8100 <212> TYPE: DNA
     8101 <213> ORGANISM: Artificial Sequence
     8103 <220> FEATURE:
     8104 <223> OTHER INFORMATION: Oligonucleotide
     8106 <400> SEQUENCE: 210
E--> 8107 accetatgtg tgatctgcct caaacccaca gcctgggtag c
     8108 (41)
8110 <210> SEQ ID NO: 211
     8111 <211> LENGTH: 46
     8112 <212> TYPE: DNA
     8113 <213> ORGANISM: Artificial Sequence
     8115 <220> FEATURE:
     8116 <223> OTHER INFORMATION: Oligonucleotide
     8118 <400> SEQUENCE: 211
E--> 8119 aaggateete atteettaet tettaaaett tettgeaagt ttgttg
     8120 46
19072 210> SEQ ID NO: 500
```

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/658,355A

19073 <211> LENGTH: 46

RAW SEQUENCE LISTING

DATE: 03/16/2004

PATENT APPLICATION: US/10/658,355A

TIME: 07:58:59

Sprins

Input Set : D:\923seq.002

Output Set: N:\CRF4\03162004\J658355A.raw

19074 <212> TYPE: DNA

19075 <213> ORGANISM: Artificial Sequence

19077 <220> FEATURE:

19078 <223> OTHER INFORMATION: primer reverse INFA-E159H

19080 <400> SEQUENCE: 500

E--> 19081 saggatecte attecttact tettaaactg tgttgeaagt ttgttg
19082 46 19084 <210> SEQ ID NO: 501

19085 <211> LENGTH: 46

19086 <212> TYPE: DNA

19087 <213> ORGANISM: Artificial Sequence

19089 <220> FEATURE:

/46

19090 <223> OTHER INFORMATION: primer reverse INFA-E159Q

19092 <400> SEQUENCE: 501

E--> 19093 aaggateete atteettaet tettaaaete tgttgeaagt ttgttg

19094

## VERIFICATION SUMMARY

DATE: 03/16/2004 TIME: 07:59:00 PATENT APPLICATION: US/10/658,355A

Input Set: D:\923seq.002

Output Set: N:\CRF4\03162004\J658355A.raw

```
L:7454 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:55 SEQ:182
L:7454 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:55 SEQ:1827
L:7466 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:55 SEQ:1837
L:7478 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:35 SEQ:1848
L:7490 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:31 SEQ:1858
L:7502 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:31 SEQ:1858
L:7514 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:18 SEQ:1868
L:7526 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:31 SEQ:1878
L:7538 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:31 SEQ:1888
L:7550 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:48 SEQ:1898
L:7562 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:48 SEQ:1908
L:7574 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:48 SEQ:19188
M:254 Repeated in SegNo=192
M:254 Repeated in SeqNo=192
L:7587 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:20 SEQ:193
 L:7599 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:21 SEQ:194
 L:7615 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:195 after pos.:0
 L:7615 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:22 SEQ:195
 L:8083 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:41 SEQ:208
 L:8095 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:46 SEQ:209
 L:8107 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:41 SEQ:210
 L:8119 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:46 SEQ:211
 L:19081 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:46 SEQ:500
 L:19093 M:254 E: No. of Bases conflict, LENGTH:Input:0 Counted:46 SEQ:501
```